

# Managing Diabetes in Long-Term Care Facilities: Benefits of Switching from Human Insulin to Insulin Analogs

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The unique requirements of residents with diabetes in long-term care (LTC) facilities necessitate a protocol-driven, individualized approach to care. Established treatment guidelines for the management of diabetes are written with the general population in mind and, although the principles remain the same in LTC patients, clinical priorities and strategies may need to be modified, and glycemic goals should be balanced against quality of life. This article identifies and explores the institutional, staff, patient and

medication-related factors that contribute to the complexity of delivering optimal diabetes care in the LTC setting, and focuses on how insulin analogs, and the pens used for their delivery, can simplify and improve care delivery while, in many cases, reducing institutional costs. (*J Am Med Dir Assoc* 2009; ■: ■-■)

**Keywords:** Long-term care (LTC); insulin analogs; insulin pens; cost

The prevalence of diabetes is high and increasing in older adults, many of whom are cared for in long-term care (LTC) residences such as nursing homes and assisted living facilities. In 2007, 23.1% of people aged 60 or older, or 12.2 million people, and about 25% of nursing home residents,<sup>1,2</sup> fulfilled the diagnostic criteria for diabetes, most with type 2 disease.<sup>3</sup> Because many patients in LTC facilities have diabetes-related complications secondary to accelerated microvascular and macrovascular disease (Table 1)<sup>4</sup> as well as other comorbidities requiring multiple medications, effective diabetes management is complex and requires a protocol-driven, team-based, individualized approach to care.

It is now widely accepted that intensive control of blood glucose that maintains HbA1c to lower than 7%, as recommended by the American Diabetes Association (ADA)<sup>5</sup> or even as low as 6.5% or less, can reduce microvascular complications,<sup>6-9</sup> and may reduce cardiovascular disease.<sup>7,8,10-12</sup> As such, there are persuasive health-related and economic reasons for treatment approaches that achieve these targets in the population as a whole. However, the recently reported long-term ADVANCE and Veterans Affairs Diabetes Trials, in patients with type 2 diabetes and increased cardiovascular risk, demonstrated no cardiovascular benefit to intensive glycemic control, whereas the ACCORD study, undertaken in a similar population, was terminated early after concluding

that intensive treatment to achieve near-normal glycemic control was associated with increased mortality in this high-risk population.<sup>13</sup> Although researchers remain unsure as to the precise mechanisms underlying the results from ACCORD, the findings support a slightly more prudent approach that brings HbA1c toward ADA target levels in high-risk adults, without the need to treat unduly aggressively. Indeed, a recent joint position statement issued by the ADA in conjunction with the American College of Cardiology Foundation and the American Heart Association in relation to ACCORD, ADVANCE, and the VA Diabetes Trial, stressed the need for an individualized approach to treatment, and less stringent HbA1c targets in patients with limited life expectancy, advanced microvascular or macrovascular complications, extensive comorbid conditions, or those with longstanding diabetes in whom the A1c goal is difficult to attain.<sup>14</sup> As such, although the principles of diabetes management are the same in older as in younger adults, national targets may not be appropriate for all LTC residents<sup>5,15,16</sup>; clinical priorities and strategies must be tailored to the individual and glycemic goals balanced against quality of life and relaxed in some patients. These caveats notwithstanding, the benefits of good glycemic control in older adults, including reduced incidence and progression of chronic complications, improved cognitive function, fewer infections, reduced incontinence, fewer emergency room visits, greater overall well-being and possibly reduced mortality rate,<sup>17,18</sup> would support a proactive approach.

The aim of this review was to identify and discuss the institutional, staff, patient, and medication-related factors that contribute to the complexity of delivering optimal diabetes care in the LTC setting, and to offer guidance on how insulin analogs, and the pens used for their delivery, can facilitate improved care delivery, while in many cases reducing institutional costs.

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**Table 1.** *Complications Associated with Diabetes in the Elderly*<sup>4</sup>

117	Confusion, acceleration of cognitive impairment
118	Decline in ability to perform activities of daily living
119	Dehydration
120	Depression
121	Excessive skin problems (infection, ulcers, delayed wound healing)
122	Eye problems (visual blurring, visual loss)
123	Falls
124	Foot ulcers, foot deformities, gangrene
125	Frequent infections
126	Increased pain perception, neuropathy
127	Nonketotic hyperosmolar coma
128	Oral health problems (caries, periodontal disease, tooth loss, dry mouth, burning mouth)
129	Recent change in weight (gain or loss)
130	Stroke
131	Urinary frequency, nocturia, urinary incontinence
132	Worsening cardiac ischemia, silent ischemia

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## 138 DIABETES MANAGEMENT IN THE LONG-TERM CARE SETTING

140 The recommended model of care for adults with diabetes  
141 in the LTC setting is summarized in the clinical practice  
142 guideline, "Diabetes management in the long-term care  
143 setting," published by the American Medical Directors  
144 Association (AMDA).<sup>4</sup> In addition to providing a stepwise  
145 diagnostic and treatment algorithm, the guideline emphasizes  
146 the need for a systematic, interdisciplinary team-based  
147 approach to care, the use of outcome and process indicators  
148 to measure a facility's performance in diabetes management,  
149 and continued monitoring of patients.

150 Although it is recognized that lifestyle changes may not  
151 always be possible, the consensus opinion of several key  
152 organizations (ADA, AMDA, and American Dietetic  
153 Association)<sup>3,4,19</sup> is that residents should be offered a regu-  
154 lar diet, physical activity should be encouraged to the ex-  
155 tent possible, and initial management of diabetes should  
156 include pharmacological treatment with oral antidiabetic  
157 agents (OADs), and metformin in particular. When an ini-  
158 tial OAD fails, combination therapy using a selection of  
159 agents with different mechanisms of action is recommen-  
160 ded, but once OADs alone are unable to achieve individ-  
161 ual glycemic goals, insulin may be indicated. Timely  
162 initiation of insulin helps bring glucose levels toward tar-  
163 get, mitigating the hyperglycemia brought on by progres-  
164 sive beta cell exhaustion and glucotoxicity, and slowing  
165 development of diabetes-related complications. Insulin  
166 can also improve overall well-being in residents with  
167 poor glycemic control and weight loss (in whom it typi-  
168 cally increases weight by 4–5 kg), and in patients whose  
169 primary symptoms are those resulting from chronic hyper-  
170 glycemia.<sup>20</sup> The use of insulins and insulin delivery devices  
171 that provide effective glycemic control while simplifying  
172 administration, and ensuring staff and patient safety, can  
173 be of particular value in LTC patients, because of the spe-  
174 cific needs of this population.

## CHALLENGES OF MANAGING DIABETES IN OLDER ADULTS

### Disease-Specific Factors

175 Multiple diabetic complications and age-related changes  
176 in metabolic processes affect drug pharmacokinetics and  
177 pharmacodynamics, thereby increasing the difficulty of  
178 managing diabetes in LTC residents, in whom the potential  
179 for, and dangers of, hypoglycemia are greater.<sup>21,22</sup> Predicting  
180 the timing of peak insulin action is a significant challenge  
181 and, furthermore, hypoglycemia is more difficult to detect  
182 in those with autonomic neuropathy or cognitive deficits  
183 such as dementia, and may present in a different way than  
184 in younger patients.<sup>23</sup> Visual and cognitive impairment,  
185 anxiety and depression, reduced manual dexterity, and  
186 irregular meal consumption common in older patients with  
187 diabetes, as well as the challenge of complex or awkward  
188 delivery systems, further increase the difficulty of self-  
189 administering and adhering to treatment in the minority of  
190 patients who choose to self-administer insulin, particularly  
191 in the assisted living setting. HbA1c goals, medications,  
192 and mode of administration must therefore be tailored to  
193 the individual, and balanced against hypoglycemic risk and  
194 overall quality of life.<sup>13</sup>

### Institutional Factors

195 The operational circumstances of many LTC facilities may  
196 themselves increase the difficulty of caring for patients with  
197 diabetes. Staff shortages, frequent staff turnover, and poor  
198 compensation make hiring and retaining qualified staff  
199 difficult. In the authors' experience, outdated practices,  
200 including the persistence of "diabetic" or "no concentrated  
201 sweet" diets, no longer recommended by national organiza-  
202 tions,<sup>4</sup> remain in many institutions. Similarly, glucose logs  
203 are not reviewed in sufficient detail or with the necessary  
204 frequency to implement logical changes to insulin or OAD  
205 regimens.<sup>24</sup> In the authors' opinion, there is often excessive re-  
206 liance on HbA1c, or on isolated fasting glucose levels for as-  
207 sessing metabolic control. Further to this, there is frequently  
208 variance in provider estimates of appropriate HbA1c targets,  
209 even for practitioners working in the same institution. For ex-  
210 ample, in a year-long physician and nurse practitioner (n = 18  
211 total) survey of care delivery practices in a nursing facility of  
212 an urban teaching hospital, 56% of providers expressed the  
213 opinion that a target HbA1c of 7% was appropriate for LTC  
214 residents, whereas 22% identified 8% and another 22% gave  
215 9% as the appropriate target.<sup>25</sup> It is also noteworthy that  
216 many facilities do not test postprandial glucose (PPG) level,  
217 known to be an independent risk factor for cardiovascular  
218 events,<sup>26</sup> possibly exposing patients to unnecessary additional  
219 cardiovascular risk. When using insulin, infrequent rotation of  
220 injection sites, and improper timing of insulin to meals, can all  
221 contribute to suboptimal care.

### Staff Factors

222 Owing to inadequate educational opportunities and time  
223 constraints, nurses and ancillary staff may have outdated or  
224 insufficient knowledge of diabetes,<sup>24</sup> resulting in  
225

233 inappropriate dosing or timing of insulin injections. Break-  
 234 down in team communication, and inadequate protocols  
 235 that do not require nursing staff to alert physicians to persis-  
 236 tently elevated blood glucose levels or marked glucose excu-  
 237 sions, deny patients the opportunity for timely improvement  
 238 in glycemic control. Similarly, when physicians make treat-  
 239 ment decisions based on HbA1c alone, without examining  
 240 glucose logs, and when their familiarity and comfort with  
 241 newer treatment, and with initiating and intensifying insulin  
 242 is limited, adherence to evidence-based algorithms is un-  
 243 likely. An attitude of “therapeutic nihilism,” in which opti-  
 244 mizing glucose control is deemed pointless, or not worth  
 245 the trouble, is not uncommon when treating older and im-  
 246 paired patients, and risks creating an environment in which  
 247 suboptimal care continues unchallenged. Indeed, even in  
 248 the absence of therapeutic nihilism, experienced nursing  
 249 home physicians manage diabetes less aggressively in patients  
 250 who are both cognitively and functionally impaired than in  
 251 those who are either functionally or cognitively impaired.  
 252 This was demonstrated in a survey that reported responses  
 253 from 255 nursing home physicians with varying levels of ex-  
 254 perience and training; survey results showed a significantly  
 255 reduced frequency of routine monitoring (eg, HbA1c, basic  
 256 chemistry) and interventions (eg, routine ophthalmology  
 257 examinations) in cognitively and functionally impaired  
 258 individuals, although the investigators did not examine the  
 259 reasons for the different management strategies.<sup>27</sup> The  
 260 results may reflect physician adherence to guidelines that  
 261 recommend individualization of care in this patient group  
 262 or, alternatively, may indicate a perceived lack of benefit,  
 263 and/or concern about greater patient inconvenience, when  
 264 more intensive treatment is used in impaired patients.

### 265 Medication and Drug Delivery Factors

267 There remain a number of practices that, although  
 268 outdated, continue to be used in some LTC facilities. These  
 269 include the persistence of now discredited sliding scale  
 270 insulin protocols,<sup>28</sup> without scheduled mealtime insulin or  
 271 rational adjustment to regimens,<sup>29</sup> the tendency to use “one  
 272 size fits all” regimens, and the continued reliance on human  
 273 insulin, delivered using vial and syringe, despite compelling  
 274 data supporting the advantages of insulin analogs delivered  
 275 using insulin pens. This is discussed in detail later in this paper.

### 276 Patient Factors

278 Despite health care provider recommendation, patients  
 279 may be reluctant to start insulin, or to accept multiple daily  
 280 doses. Reasons for this “psychological insulin resistance” in  
 281 the general adult population are discussed at length  
 282 elsewhere<sup>30–32</sup> and may include feelings of defeat or personal  
 283 failure regarding disease management, fear of hypoglycemia  
 284 or weight gain, needle anxiety, and concern about being  
 285 able to cope with what seems a complicated new treatment.  
 286 Although there are no studies addressing this issue specifically  
 287 in the elderly, clinical experience suggests that concerns  
 288 about adverse effects and the difficulties of starting a new  
 289 and potentially complex treatment late in life would  
 290 predominate. LTC subjects who do start insulin may be

frustrated in their efforts by complications resulting from  
 their diabetes and/or concurrent disease processes; for  
 example, fasting hypoglycemia followed by very high blood  
 glucose levels later in the day may combine to make  
 treatment feel overwhelming. The greater potential for  
 adverse effects and drug interactions in elderly patients neces-  
 sitate caution when prescribing, and greater vigilance in  
 monitoring adverse effects, particularly since hepatic and  
 renal impairment are relatively common in older residents  
 with diabetes.

### “State of the Field” Factors

The lack of studies evaluating optimal glycemic and  
 HbA1C goals, as well as lipid and blood pressure targets, in  
 elders and nursing home patients leaves health care providers  
 with little evidence-based guidance.<sup>33</sup> Likewise, there is often  
 insufficient attention to evidence that indicates practices to  
 be avoided.<sup>33</sup> Individual facilities frequently lack protocols  
 for monitoring diabetes or assessing facility-wide diabetes  
 management, resulting in inconsistencies in treatment and  
 different approaches to management even within institutions.  
 Health care providers are therefore urged to consult guidelines  
 designed with the long-term patient population in mind, all of  
 which have been prepared by expert consensus and analysis of  
 available evidence. These include the AMDA clinical  
 practice guidelines<sup>4</sup> as well as those from the American  
 Geriatrics Society in collaboration with the California  
 Healthcare Foundation,<sup>16</sup> and those by Zarowitz et al.<sup>34</sup>

### THE ECONOMIC BENEFITS OF OPTIMIZING GLYCEMIC CONTROL

In addition to improving patient well-being and prognosis,  
 there are compelling economic reasons to strive for optimal  
 glycemic control, and particularly for reduction of chronic  
 complications. The estimated cost of diabetes in the United  
 States in 2007 was \$174 billion, of which \$58 billion was  
 used to treat chronic complications.<sup>35</sup> In a 2002 retrospective  
 database analysis of more than 3000 health maintenance  
 organization claims, patients with type 2 diabetes whose  
 HbA1c was consistently greater than 7% over 1 year accrued  
 diabetes-related costs 32% higher than those whose HbA1c  
 remained 7% or lower (\$1540 versus \$1171),<sup>36</sup> whereas  
 a retrospective US-based analysis of more than 9000  
 commercial health plan claims in 2006, found that 12-month  
 costs for patients with diabetes and macrovascular disease  
 were more than 3 times those with diabetes but no  
 macrovascular disease (\$10,450 versus \$3,385).<sup>37</sup> Similarly,  
 a large European study (CODE -2), found that the presence  
 of both microvascular and macrovascular disease increased  
 the total cost of management by as much as 250% per  
 patient.<sup>38</sup> A systematic review evaluating economic and  
 resource utilization in US-managed care organizations found  
 that pharmacy costs were primarily driven by medications  
 used to treat diabetic complications, with only 30% expended  
 on glycemic control.<sup>39</sup> In this study, improving and maintain-  
 ing glycemic control led to overall cost savings, irrespective of  
 the medications used. Interestingly, the higher initial  
 pharmacy costs associated with starting insulin or using

349 newer, more expensive, insulin formulations were more than  
350 offset by reduced inpatient hospital admissions, sometimes  
351 within less than 1 year.

352 Although data are lacking on the cost of complications  
353 specifically in the LTC setting, it is clear that they  
354 significantly increase the cost of care. Strategies to guide  
355 the safe attainment of glycemic control not only ensure  
356 high standards of patient care but also contain costs by  
357 forestalling the development of, or worsening of, chronic  
358 diabetic complications, reducing the incidence of hypoglyce-  
359 mic events, reducing needlestick injuries to staff and patients,  
360 and by limiting the frequency of hospitalization and  
361 emergency room visits.

## 363 INSULIN ANALOGS AND DELIVERY PENS CAN 364 IMPROVE PATIENT CARE AND REDUCE 365 INSTITUTIONAL COSTS

### 367 Clinical Benefits of Insulin Analogs

368 Insulin analogs, developed by making minor amino acid  
369 substitutions to the human insulin molecule, have pharmaco-  
370 kinetic and pharmacodynamic profiles that closely approxi-  
371 mate those of endogenous insulin, resulting in a more  
372 consistent glycemic effect than human insulins; similar, or  
373 even lower, HbA1c; better control of postprandial and fasting  
374 glucose; and reduced rates of hypoglycemia, particularly at  
375 night (Table 2).<sup>40</sup> Retail prescriptions for insulin analogs  
376 among nonhospitalized patients now far exceed those for  
377 human insulins (approximately 48,000 versus 19,000  
378 units  $\times$  10<sup>6</sup> in 2007), but LTC formularies generally do not  
379 reflect this preference: 2007 data for prescriptions in LTC  
380 facilities indicate comparable prescription data of 3100 and  
381 3500 units  $\times$  10<sup>6</sup> for insulin analogs and human insulins,  
382 respectively (IMS Health; data on file 2007).

383 Data on basal insulins, usually the first choice for  
384 once-daily treatment, demonstrate several advantages for  
385 analogs, particularly a reduced incidence of hypoglycemia.  
386 In a meta-analysis comparing insulin detemir (2 studies,  
387 578 patients) and insulin glargine (6 studies, 1715 patients)  
388 with neutral protamine Hagedorn (NPH) insulin, in studies  
389 of 24 to 52 weeks' duration, patients were found to have  
390 a similar degree of metabolic control, using HbA1c as  
391 a surrogate end point, as well as similar rates of severe  
392 hypoglycemia and overall adverse events, but the incidences  
393 of overall, symptomatic, and nocturnal hypoglycemia were all  
394 statistically significantly reduced in the insulin analog  
395 groups.<sup>41</sup> Similarly, a systematic review of 6 head-to-head  
396 insulin glargine versus NPH studies found equivalence with  
397 respect to reduction in HbA1c, but patients receiving  
398 glargine experienced equivalent or lower fasting plasma  
399 glucose (FPG) levels and less frequent nocturnal hypoglyce-  
400 mia.<sup>42</sup> Additional data suggest that this reduction in  
401 nocturnal hypoglycemia holds true only during once-daily  
402 administration of NPH; during twice-daily dosing, the  
403 superiority of glargine may no longer be apparent.<sup>43</sup>

404 Individual trial data highlight some of the advantages of  
405 basal insulin analogs in patients with type 2 diabetes. In a 2-  
406 week treat-to-target study of 498 insulin-naïve type 2

**Table 2.** *Advantages of Insulin Analogs over Human Insulin in the Long-Term Care Setting*

More consistent, predictable glycemic effect	407
Equivalent or lower HbA1c	408
Better control of fasting and postprandial glucose levels	409
Reduced risk of hypoglycemia, particularly at night	410
Weight-sparing effect	411
Insulin delivery using pens improves dosing accuracy, reduces hypoglycemia and is preferred by patients over vial/syringe	412
Insulin pens reduce needlestick injuries and dosing errors	413
Replacement of vial/syringe with insulin pens results in significant all-cause and hypoglycemia-related cost savings	414

421 patients poorly controlled on OADs, patients taking  
422 once-daily insulin detemir in the evening experienced  
423 reductions in 24-hour and nocturnal hypoglycemia that were  
424 53% ( $P = .02$ ) and 63% ( $P = .03$ ) lower, respectively, than  
425 those in patients taking evening NPH insulin.<sup>44</sup> When insulin  
426 detemir was given in the morning, the incidence of nocturnal  
427 hypoglycemia fell by 87% compared with that observed in pa-  
428 tients taking evening NPH insulin ( $P < .001$ ). Similarly, addi-  
429 tion of insulin glargine to failing OADs in a group of 756  
430 overweight individuals allowed approximately 60% to reach  
431 HbA1c 7.0% or less, with significantly fewer patients on glar-  
432 gine experiencing nocturnal hypoglycemia (33% versus 27%,  
433  $P < .05$ ).<sup>45</sup> In a post hoc analysis of 3 open-label multinational  
434 phase III studies in which older ( $\geq 65$  years) and younger (18–  
435 64 years) individuals with type 2 diabetes were treated for 22 to  
436 26 weeks with basal insulin (NPH insulin or insulin detemir)  
437 plus mealtime insulin or OADs, control of HbA1c and FPG  
438 were equivalent in the 416 older adults receiving insulin dete-  
439 mir or NPH insulin, but the relative risk of hypoglycemia fa-  
440 vored insulin detemir (RR = 0.59), as did the reduced  
441 weight gain (mean treatment difference  $-1.02$  kg).<sup>46</sup> When  
442 choosing an initial insulin regimen in type 2 patients inade-  
443 quately controlled on OADs, studies show that the addition  
444 of a basal analog to OADs offers greater efficacy with less hypo-  
445 glycemia than starting twice-daily human premix without  
446 OADs; this was true for both the general adult population<sup>47</sup>  
447 and for older adults.<sup>48</sup>

448 Observational study results are a valuable addition to  
449 clinical trial data in that they typically reflect real-life  
450 practice and outcomes. The very large Predictable Results  
451 and Experience in Diabetes through Intensification and  
452 Control to Target: an International Variability Evaluation  
453 (PREDICTIVE) trial, a multinational, prospective, observa-  
454 tional study of insulin detemir in clinical practice,  
455 demonstrated the ability of insulin detemir, with or without  
456 OADs, to significantly reduce mean HbA1c ( $-1.3\%$ ;  
457  $P < .0001$ ) and fasting glucose ( $-3.7$  mmol/L;  $P < .0001$ ),  
458 typically with once-daily injection.<sup>49</sup> There was a significantly  
459 reduced risk of hypoglycemia in patients who switched to  
460 insulin detemir from NPH insulin as well as in those starting  
461 insulin detemir after OAD failure.<sup>50</sup> PREDICTIVE also  
462 highlighted the weight-sparing effect of insulin detemir.<sup>50</sup>  
463 Similarly, in a retrospective observational study in 397  
464 patients with type 2 diabetes who switched from an

465 NPH-based to a glargine-based regimen (approximately 80%  
 466 of whom used a basal-bolus regimen before and after the  
 467 switch), HbA1c decreased by 0.31% after 1 year ( $P < .001$ )  
 468 with no significant change in weight or total daily insulin  
 469 dose.<sup>51</sup> These benefits can be maintained beyond 1 year, as  
 470 demonstrated in a large 9-month observational study, with  
 471 a 20-month extension period, in which glargine was added  
 472 to the treatment regimen in 12,216 patients (mean  
 473 age =  $63.9 \pm 11.3$  years) inadequately controlled on  
 474 OADs.<sup>52</sup> The addition of glargine significantly reduced  
 475 HbA1c (by 1.5%) and fasting blood glucose (by 69 mg/dL)  
 476 after 3 months, an improvement that was maintained at  
 477 9 months, and indeed at 20 months in 2721 patients who con-  
 478 tinued treatment during the extension. These benefits were  
 479 achieved with no increase in body mass index. According  
 480 to a comprehensive evaluation of basal insulin analogs, the  
 481 addition of a once-daily long-acting insulin analog to  
 482 OADs typically lowers HbA1c by about 1.5% over a period  
 483 of 20 to 24 weeks, with reduced hypoglycemia at equivalent  
 484 levels of glycemic control.<sup>53</sup> Data are fairly consistent for  
 485 patients with baseline HbA1c of 8.5% or lower; for those  
 486 with HbA1c in excess of this, a basal-only approach is  
 487 probably insufficient to bring glycemic control to target,  
 488 although, as previously explained, this may be less essential  
 489 in the LTC setting.

490 Owing to the progressive nature of type 2 diabetes, many pa-  
 491 tients will ultimately need both basal and mealtime (prandial)  
 492 insulin supplementation. This can be achieved simply in well-  
 493 controlled patients with a consistent eating pattern by using  
 494 once- or twice-daily premixed insulin; patients who are poorly  
 495 controlled may be considered candidates for basal-bolus ther-  
 496 apy.<sup>4</sup> Premixed insulin analogs offer similar HbA1c reduction  
 497 to human premixes,<sup>54,55</sup> but by allowing flexible injection tim-  
 498 ing relative to meals, offer greater treatment flexibility, a par-  
 499 ticular advantage in the LTC population whose meal  
 500 consumption is often inconsistent or whose meals may be de-  
 501 layed by factors beyond their control. When compared with  
 502 long-acting insulins, premixed analogs offer significantly bet-  
 503 ter postprandial control because of inclusion of the short-act-  
 504 ing mealtime insulin component.<sup>54</sup> In the large PRESENT  
 505 study, a multinational observational study of patients poorly  
 506 controlled on human insulin  $\pm$  OADs or OAD therapy alone,  
 507 switching from biphasic human insulin to biphasic insulin as-  
 508 part 70/30 significantly reduced mean HbA1c ( $-1.6\%$ ), FPG  
 509 ( $-52.6$  mg/dL), and PPG ( $-86.4$  mg/dL).<sup>56</sup> The rate of  
 510 overall, major, and nocturnal hypoglycemia fell from 8.9 to  
 511 2.2 episodes/patient year, 0.7 to 0.1 episodes/patient year,  
 512 and 2.9 to 0.5 episodes/patient year, respectively. Premixed in-  
 513 sulin analogs are available in prefilled pens, and allow for  
 514 twice-daily injection.

515 Patients inadequately controlled on a premix may benefit  
 516 from intensification of therapy using analog-based basal-bolus  
 517 therapy. Study data indicate that a switch from a premix to an  
 518 insulin glargine-based regimen, either in combination with  
 519 OADs or as the basal component of a basal-bolus regimen,  
 520 can improve glycemic control as well as tolerability and  
 521 treatment satisfaction.<sup>57,58</sup> In a retrospective observational  
 522 analysis of 345 patients with type 2 diabetes, of whom 48%

were using premixed insulin only and 38% premixed  
 insulin + OAD, a switch to insulin glargine  $\pm$  OADs/  
 prandial insulin decreased HbA1c by 0.53% after 12 months  
 ( $P < .001$ ) with no change in weight.<sup>59</sup> Not surprisingly,  
 there was a significant increase in the number of patients  
 who needed to add prandial insulin (from 16.2% to 73.9%;  
 $P < .001$ ) and an increase in total mean insulin administra-  
 tion ( $P < .001$ ).

Candidates for basal-bolus therapy are best managed using  
 rapid-acting analogs as the mealtime insulin. Although  
 relevant clinical trials of basal-bolus therapy in type 2  
 diabetes are limited, a 26-week multinational study in  
 patients with type 2 diabetes (mean age = 60 years), random-  
 ized to either insulin detemir or NPH insulin, both with  
 mealtime insulin aspart, found equivalent HbA1c levels at  
 study end point, similar FPG reductions, and equivalent  
 hypoglycemic risk.<sup>60</sup> However, detemir was associated with  
 reduced intrasubject daily variation in fasting self-monitored  
 blood glucose, and significantly less weight gain (1.0 versus  
 1.8 kg;  $P = .02$ ). Similarly, a 22-week study of 395 individuals  
 with type 2 diabetes found that an all-analog insulin regimen  
 (insulin detemir plus insulin aspart) was associated with mean  
 weight gain of 0.5 kg versus 1.1 kg in patients on an  
 all-human insulin regimen (NPH insulin plus regular human  
 insulin) ( $P < .05$ ).<sup>61</sup>

In addition to the benefits of insulin analogs themselves,  
 the delivery devices used for injection offer numerous  
 advantages for patients and staff. Patients typically find  
 insulin pens simple to use<sup>62,63</sup> and compared with syringe/  
 vial, experience fewer hypoglycemic episodes<sup>64</sup> and express  
 an overall preference,<sup>65,66</sup> resulting in greater adherence to  
 treatment and improved quality of life. This is of particular  
 value in subacute care patients who can be taught how to  
 self-inject before returning home.

### Economic Benefits of Insulin Analogs and Delivery Pens

By virtue of greater convenience and improved safety  
 compared with vial and syringe,<sup>67</sup> insulin pens can reduce  
 hypoglycemia rates, increase treatment satisfaction, and  
 facilitate adherence, often at an overall cost saving.<sup>66,68</sup> The  
 use of pens also reduces emergency visits and hospitaliza-  
 tion.<sup>64,66</sup> For example, in one database analysis of managed-  
 care claims for more than 40 million covered lives, adults  
 with type 2 diabetes who switched from using vial/syringe to  
 deliver human or analog insulin to using a prefilled insulin  
 analog pen demonstrated significantly improved treatment  
 adherence (increased from 62% to 69%;  $P < .01$ ), reduced  
 hypoglycemic risk (odds ratio [OR] = 0.5; 95% confidence  
 interval [CI]: 0.37, 0.68;  $P < .05$ ), and significant reductions  
 in emergency room and office visits ( $P < .05$  for both).<sup>66</sup>  
 This led to all-cause and hypoglycemia-attributable  
 treatment savings of \$1590 per patient ( $P < .01$ ), and \$788/  
 patient ( $P < .01$ ), respectively. In a second database study of  
 486 privately insured patients converting from human or  
 analog insulin delivered by vial/syringe to an insulin analog  
 pen, the incidence of hypoglycemia again decreased signifi-  
 cantly (OR = 0.4; 95% CI: 0.27, 0.61;  $P < .05$ ) as did

hypoglycemia-related emergency room and physician office visits ( $P < .05$  for both); total annual and hypoglycemia-attributable treatment costs fell by \$1748/patient and \$908/patient, respectively.<sup>64</sup> There are also data suggesting that treatment costs can be reduced in Medicaid patients poorly controlled on OADs, by initiating an insulin analog pen rather than starting on vial/syringe when insulin becomes necessary.<sup>69</sup> In this large database analysis of Medicaid patients, total annualized health care costs were significantly lower in patients starting insulin using a pen as opposed to vial/syringe (\$14,857 versus \$31,765), as were hospital, diabetes-related, and outpatient costs ( $P < .05$  for all).<sup>69</sup>

Improving the safety and accuracy of insulin delivery and blood glucose monitoring are important aspects of care. The Centers for Disease Control and Prevention (CDC) offer specific guidance on measures to reduce the risk of blood-borne transmission of pathogens during these procedures.<sup>70</sup> To this end, insulin pens, some used with dedicated needles that include safety features such as automatic safety locks (eg, NovoFine Autocover 8-mm needle [Novo Nordisk A/S, Bagsvaerd, Denmark], and the BD Autosheild Pen Needle [Becton Dickinson, Franklin Lakes, NJ]), can help address the relatively high rate of staff needlestick injuries resulting from use of disposable syringes (448 per 1000 nurses in one recent study),<sup>71</sup> and the associated risks and cost of viral hepatitis infection and missed days of work. The benefits of prefilled insulin pens include their use for delivery of premixed insulins, thereby avoiding mixing errors when withdrawing insulin from multiple vials, as well as the possibility of leaving them in a patient's personal medication drawer to avoid multiple user errors with vial/syringe, or selection of the wrong insulin; color coding may also help in selection of the appropriate pen and insulin. From an institutional perspective, the formulary simplification resulting from a complete switch from conventional insulins with vial and syringe, to insulin analogs and pens, can reduce the frequency and cost of dosing errors.<sup>72</sup>

## CONCLUSIONS

Delivering high-quality care to LTC residents with diabetes requires an understanding of the unique needs of this population, and of the complexities involved in treating older individuals with multiple comorbidities in an institutional setting. There is a pressing need for more clinical trials that focus on elderly patients with diabetes, as well as improved treatment guidelines and algorithms for this population. Results of such research will help LTC health care providers manage diabetes in an evidence-based manner, as well as raising treatment expectations.

Although there are many challenges to delivering care, those that result from medication-related factors can often be met by switching from human to analog insulins, and by using insulin pens in selected patients. Insulin analogs generally provide more consistent glycemic control than human insulins, equivalent or improved HbA1c, better control of PPG and fasting glucose, and reduced rates of hypoglycemia, without the problem of weight gain. When used with insulin pens, which patients who self-inject

generally prefer and find easy to use, these agents address several of the widely identified barriers to insulin use and facilitate treatment adherence. From an institutional perspective, these clinical benefits translate into economic advantages and significant cost savings over time. As the aging population of patients with diabetes increases in number, switching from human to analog insulins, and from vial/syringe to insulin pens, will provide clinical, safety, and economic benefits to both patients and LTC facilities.

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